

Form PTO-1449 (modified)

AUG 08 2001

List of Patents and Publications for Applicants

## INFORMATION DISCLOSURE STATEMENT

(Use several sheets if necessary)

Atty. Docket No.

ARCD:374US/GNS

Serial No.

09/835,082

Applicant

Mark J. Ratain, Federico Innocenti and Lalitha Iyer

Filing Date:

April 12, 2001

Group:

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## U.S. Patent Documents

Exam. Init.	Ref. Des.	Document Number	Date	Name	Class	Sub Class	Filing Date of App.

## Foreign Patent Documents

Exam. Init.	Ref. Des.	Document Number	Date	Country	Class	Sub Class	Translation Yes/No

## Other Art (Including Author, Title, Date Pertinent Pages, Etc.)

Exam. Init.	Ref. Des.	Citation
CW	C1	Bible, et al., "Cytotoxic synergy between flavopiridol (NSSC 649890, L86-8275) and various antineoplastic agents: the importance of sequence of administration," <i>Cancer Res.</i> , 57:3375-3380, 1997.
	C2	Bible and Kaufmann, "Flavopiridol: acytotoxic flavone that induces cell death in noncycling A549 human lung carcinoma cells," <i>Cancer Res.</i> , 56:4856-4861, 1996.
	C3	Bock et. al., In: Conjugation reactions in biotransformation, Elsevier, North Holland Biomedical Press, p. 357-364, 1978.
	C4	Carlson, et al., "Flavopiridol induces G <sub>1</sub> arrest with inhibition of cyclin-dependent kinase (CDK) 2 and CDK4 in human breast carcinoma cells," <i>Cancer Res.</i> , 56:2973-2978, 1996.
	C5	Cascorbi et al., "Frequency of single nucleotide polymorphisms in the p-glycoprotein drug transporter MDR1 gene in white subjects," <i>Clinic. Pharmacol Ther.</i> , 69:169-174, 2001.
	C6	Chien et al., "In vitro evaluatino of flavopiridol, a novel cell cycle inhibitor, in bladder cancer," <i>Cancer Chemother Pharmacol</i> , 44:81-87, 1999.
	C7	Coffman, et al. "The glucuronidation of opioids, other xenobiotics and androgens by human ugt2b7y(268) and ugt2by7h(268)," <i>Drug Metab Dispos</i> , 26:73-77, 1998.
	C8	Czech, et al., "Antitumoral activity of flavone L 86-8275," <i>nt J Oncol.</i> , 6:31-66, 1995.
	C9	Decleves et al., "A new polymorphism (N21D) in the exon 2 of the human MDR1 gene enclosing the P-glycoprotein," <i>Human Mutation</i> , 15: 486, 2000.

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Cynthia M. Alder

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Exam. Init.	Ref. Des.	Citation
CW	C10	Di Carlo et al., "Flavonoids: old and new aspects of a class of natural therapeutic drugs," <i>Life Sci.</i> , 65:337-353, 1999.
	C11	Diasio et al., "Clinical pharmacology of 5-fluorouracil," <i>Clin Pharmacokinet</i> , 16:215-237, 1989.
	C12	Drees et al., "Flavopiridol (L86-8275): selective antitumor activity in vitro and activity in vivo for prostate carcinoma cells," <i>Clin Cancer Res</i> , 3:273-9, 1997.
	C13	Gutmann et al., "Modulation of multidrug resistance protein expression in porcine brain capillary endothelial cells in vitro," <i>Drug Metab Dispos.</i> 27:937-941, 1999.
	C14	Hoffmeyer et al., "Functional polymorphisms of the human multidrug-resistance gene: multiple sequence variations and correlation of one allele with p-glycoprotein expression and activity in vivo," <i>PNAS</i> , 28:97(7), 3473-3478, 2000.
	C15	Hooijberg et al., "Potent interaction of flavopiridol with MRP1," <i>British J. of Cancer</i> , 81:269-276, 1999.
	C16	Innocenti et al., "Flavopiridol metabolism in cancer patients is associated with the occurrence of diarrhea," <i>Clinical Cancer Research</i> , 6:3400-3405, 2000.
	C17	Ito et al., "Polymorphism of the abc transporter genes mdr1, mrp1 and mrp2/cmoat, in healthy japanese subjects," <i>Pharmacogenetics</i> , 11:175-184, 2001.
	C18	Iyer, et al. "Pharmacogenetics and cancer chemotherapy," <i>Eur J Cancer</i> , 34:1493-1499, 1998.
	C19	Iyer, "Inherited variations in drug-metabolizing enzymes: significance in clinical oncology," <i>Mol Diagnosis</i> , 4:327-333, 1999.
	C20	Jager et al., "Metabolism of the anticancer drug flavopiridol, a new inhibitor of cyclin dependent kinases in rat liver," <i>Life Sci.</i> , 62:1861-73, 1998.
	C21	Kusuhara, et al., "Reduced folate derivatives are endogenous substrates for cmoat in rats," <i>Am J Physiol.</i> , 275(4 Pt 1):G789-G796, 1998.
	C22	Lennard, "The clinical pharmacology of 6-mercaptopurine," <i>Eur J Clin Pharmacol.</i> , 43:329-339, 1992.
	C23	Levesque et al., "Isolation and characterization of UGT2B15(Y85): a udp-glucuronosyltransferase encoded by a polymorphic gene," <i>Pharmacogenetics</i> 7:317-325, 1997.

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<i>Qu</i>	C24	Levesque et al., "Characterization and substrate specificity of UGT2B4 (E <sup>458</sup> : a udp-glucuronosyltransferase encoded by a polymorphic gene," <i>Pharmacogenetics</i> , 9:207-16, 1999.
	C25	Lomri et al., "Hepatocellular transport: role of atp-binding cassette proteins," <i>Semin. Liv. Dis.</i> , 16: 201-210, 1996.
	C26	Losiewicz, et al., "Potent inhibition of cdc2 kinase activity by the flavonoid L86-8275," <i>Biochem Biophys Res Commun.</i> , 201:589-595, 1994.
	C27	Meech et al., "Determinants of udp glucuronosyltransferase membrane association and residency in the endoplasmic reticulum," <i>Arch Biochem Biophys.</i> , 356:77-85, 1998.
	C28	Miners, et al., "Drug glucuronidation in humans," <i>Pharmacol Ther.</i> , 51:347-369, 1991.
	C29	Muller et al., "ATP-dependent transport of amphiphilic cations across the hepatocyte canalicular membrane mediated by mdr1 p-glycoprotein," <i>FEBS Lett.</i> , 343:168-172, 1994.
	C30	Nebert, "Pharmacogenetics and pharmacogenomics: why is this relevant to the clinical geneticist?" <i>Clin Gen.</i> , 56:247-258, 1999.
	C31	Perdu and Germain, "Identification of novel polymorphisms in the pm5 and mrp1(abcc1) genes at locus 16p13.1 and exclusion of both genes as responsible for pseudoxanthoma elasticum," <i>Human Mutation</i> , 17:74-75, 2001.
	C32	Ramírez et al., "In vitro glucuronidation of flavopiridol (nsc649890) (flavo) by human liver microsomes," <i>Clin Pharmacol Ther.</i> , (abstract) 63:149, 1998.
	C33	Ratain et al., "Paradoxical relationship between acetylator phenotype and amonafide toxicity," <i>Clin. Pharmacol. Ther.</i> , 50:573-579, 1991.
	C34	Robey et al., "Overexpression of the atp-binding cassette half-transporter, abcg2 (mxr/bcrp/abcp1), in flavopiridol-resistnat human breast cancer cells," <i>Clinical Cancer Res.</i> , 7:145-152, 2001.
	C35	Rund et al, "A mutation in the promoter of the multidrug resistance gene (mdr1) in human hematological malignancies may contribute to the patogenesis of resistant disease," <i>Adv. Exp Med Biol.</i> , 457:71-75, 1999.
	C36	Sausville et al., "Cyclin-dependent kinases: initial approaches to exploit a novel therapeutic target," <i>Pharmacol Ther.</i> , 82:285-292, 1999.

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Ce	C37	Schrump et al., "Flavopiridol mediates cell cycle arrest and apoptosis in esophageal cancer cells," <i>Clin Cancer Res.</i> , 4:2885-2890, 1998.
	C38	Senderowicz et al., "Phase I trial of continuous infusion flavopiridol, a novel cyclin-dependent kinase inhibitor, in patients with refractory neoplasms," <i>J Clin Oncol.</i> , 16:2986-2999, 1998.
	C39	Shapiro et al., "Flavopiridol induces cell cycle arrest and p53-independent apoptosis in non-small cell lung cancer cell lines," <i>Clin. Cancer Res.</i> , 5:2925-2938, 1999.
	C40	Sherr, "Cancer cell cycles," <i>Science</i> , 274:1672-1677, 1996.
	C41	Stadler et al., "Flavopiridol, a novel cyclin-dependent kinase inhibitor, in metastatic renal cancer: a university of chicago phase II consortium study," <i>J Clin Oncol.</i> , 18:371-375, 2000.
	C42	Thomas et al., "Phase I clinical an dpharmacokinetic trial of flavopiridol," <i>Proc Am Assoc Cancer Res.</i> , (abstract) 38:1496, 1997.
	C43	Vezmar et al., "reversal of mrp-mediated doxorubicin resistance with quinoline-based drugs," <i>Biochem Pharmacol.</i> , 59:1245-1252, 2000.
	C44	Worland et al., "Alteration of the phosphorylation state of p34cdc2 kinase by the flavone L86-8275 in breast carcinoma cells," <i>Biochem Pharmacol.</i> , 46:1831-1840, 1993.

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